Amendments to the Claims

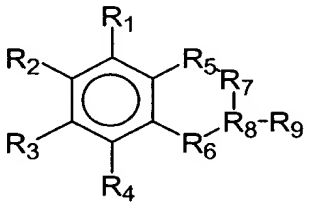
The following listing of claims will replace all prior versions, and listings, of claims in the application:

1-6. (Canceled)

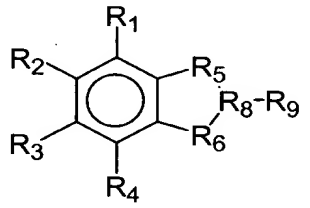
7. (Currently amended) A composition comprising an angiogenesis inhibiting compound and an anti-inflammatory drug, wherein the angiogenesis inhibiting compound is:

(1) a compound of the formula

A)

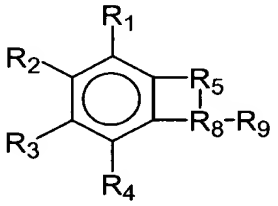


B)

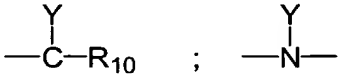


or

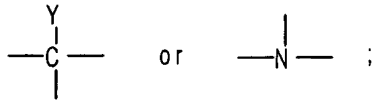
C)



wherein R_1-R_4 are each independently -H; -OH; =O; straight or branched chain alkane, alkene, or alkyne; cyclic alkane, alkene, or alkynes; a combination of cyclic and acyclic alkanes, alkenes, or alkynes; alcohol, aldehyde, ketone, carboxylic acid, ester, or ether moieties in combination with acyclic, cyclic, or combination acyclic/cyclic moieties; aza; amino, $-XO_n$ or $-O-XO_n$, where X=N and n=2, X=S and n=2 or 3, or X=P and n=1-3; or halogen; R_5-R_7 are each independently

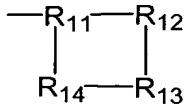


or -O-, where Y is absent and R_{10} is =O, or Y and R_{10} are each independently -H; -OH; =O; straight or branched chain alkane, alkene, or alkyne; cyclic alkane, alkene, or alkynes; a combination of cyclic and acyclic alkanes, alkenes, or alkynes; alcohol, aldehyde, ketone, carboxylic acid, ester, or ether moieties in combination with acyclic, cyclic, or combination acyclic/cyclic moieties; aza; amino, $-XO_n$ or $-O-XO_n$, where X=N and n=2, X=S and n=2 or 3, or X=P and n=1-3; or halogen; R_8 is:



and R_9 is

D)



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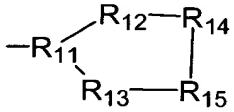
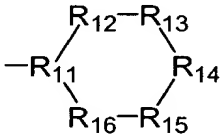


F)



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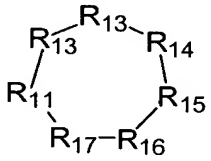
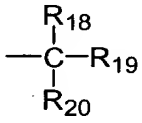
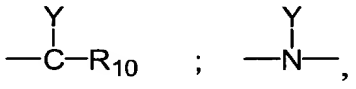


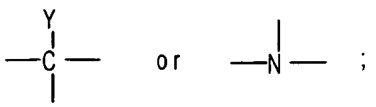
or H)



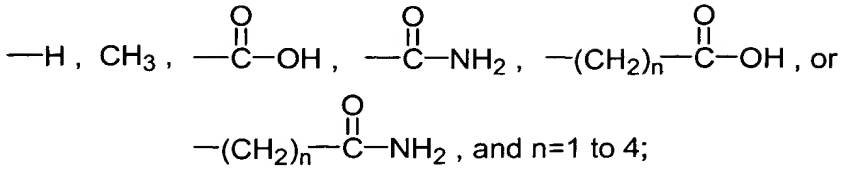
wherein each of R_{12}-R_{17} is independently



wherein R_{11} is

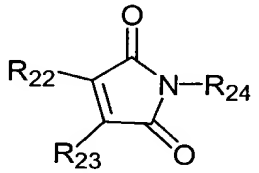


and wherein R_{18}, R_{19} and R_{20} are each independently



with the proviso that the angiogenesis inhibiting compound is not thalidomide;

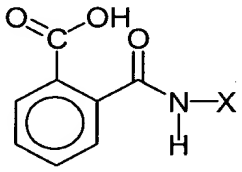
(2) a compound of the formula



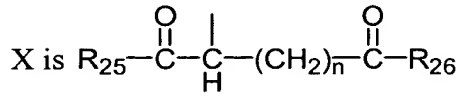
where R_{22} and R_{23} are each independently H, F, Cl, Br, I, CH_3, or $-CH_2-CH_3$; and R_{24} is H, CH_3, or $-CH_2-CH_3$;

or

(3) a compound of the formula



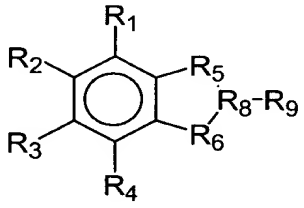
where X is R_6 as defined in (1) above, or



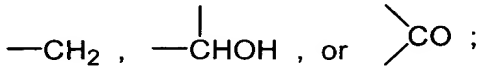
and R_{25} and R_{26} are independently -OH, -H, or $-NH_2$, and n = 1 through 4.

8. (Currently amended) The ~~angiogenesis inhibitory~~ composition of Claim 7 wherein the angiogenesis inhibiting compound is of the formula

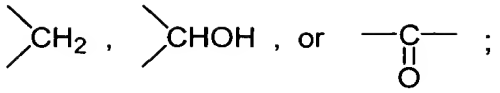
B)



and R_5 and R_6 are independently



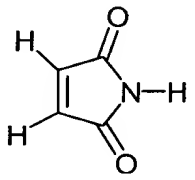
and R_9 is F) or H) wherein R_{14} and R_{16} are each independently



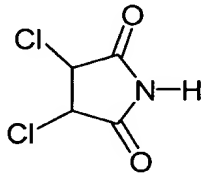
and R_{15} is -O- or  , where R_{21} is H, CH_3, or OH.

9. (Previously presented) The composition of claim 7 wherein the angiogenesis inhibiting compound is

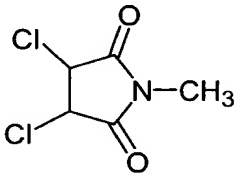
I)



J)



K)



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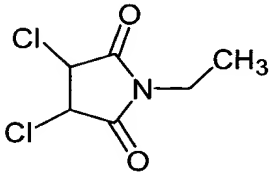
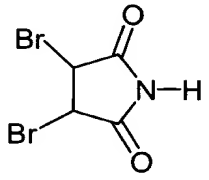
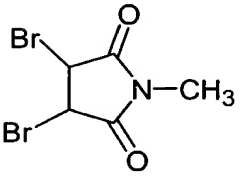


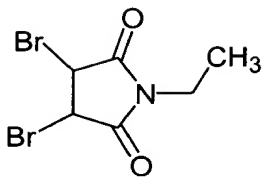
M)



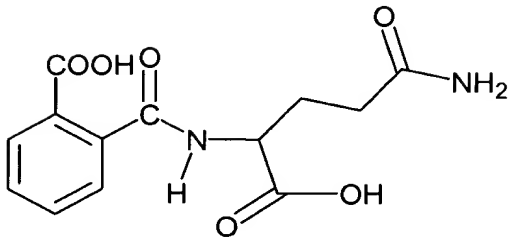
N)



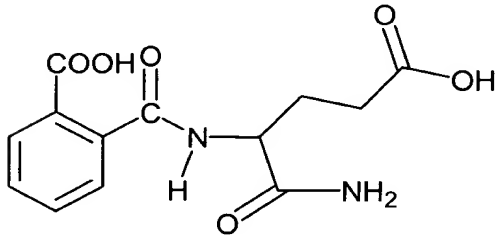
O)



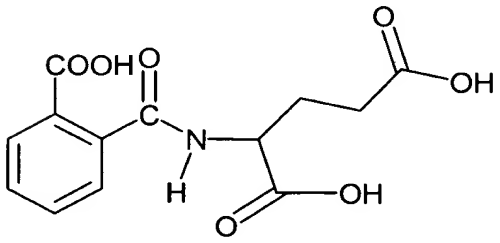
P)



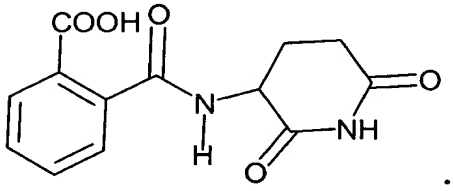
Q)



R)

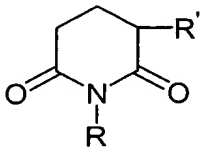


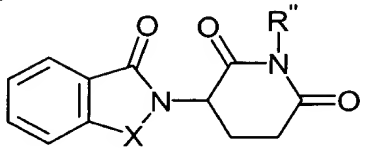
or S)



10. (Previously presented) The composition of Claim 7 wherein the angiogenesis inhibiting compound is a metabolite of thalidomide, thalidomide analog, epoxide of thalidomide, hydrolysis product thereof, hydrolysis product of thalidomide, EM-12, metabolite of EM-12, epoxide of EM-12, hydrolysis product thereof, EM-138, metabolite of EM-138, epoxide of EM-138, hydrolysis product thereof, N-phthaloyl-DL-glutamic acid, N-phtaloyl-DL-glutamine anhydride, or mixture thereof.

11. (Previously presented) The composition of Claim 10 wherein the angiogenesis inhibiting compound is

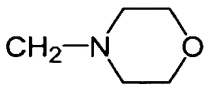
(I) 

or (II) 

wherein

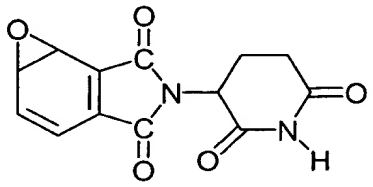
R is H, (C_1-C_6)alkyl, phenyl, or benzyl; and R' is phthalimido or succinimido; wherein

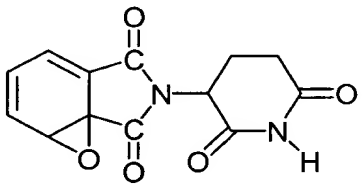
X is CH_2 or C=O; and R" is H, -CH_2CH_3, -C_6H_5, -$CH_2C_6H_5$, -$CH_2CH{=}CH_2$, or

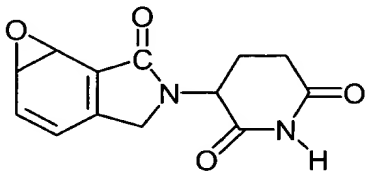


or (III) hydrolysis products of (II), wherein R" is H and the piperidino ring or both the piperidino and the imido ring are hydrolyzed.

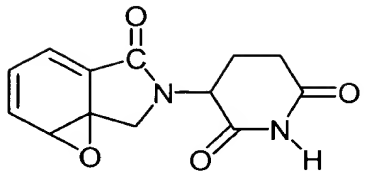
12. (Previously presented) The composition of Claim 10 wherein the angiogenesis inhibiting compound is

III) 

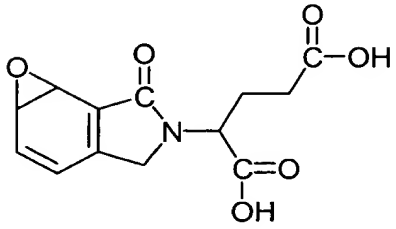
IV) 

V) 

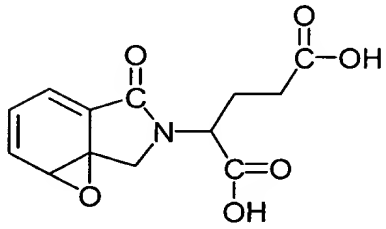
VI)



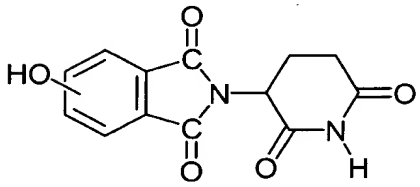
VII)



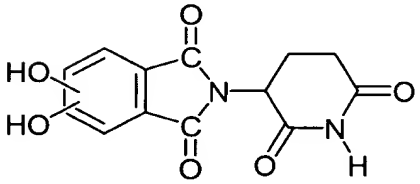
VIII)



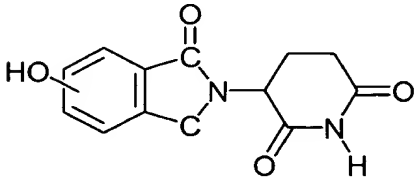
IX)



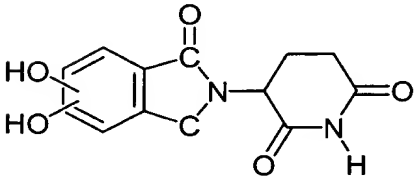
X)



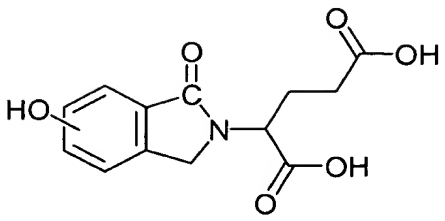
XI)



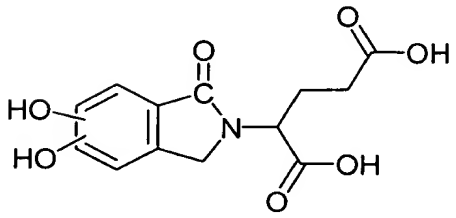
XII)



XIII)



or XIV)



13-20. (Canceled).

21. (Previously presented) The composition of Claim 7 wherein the antiinflammatory drug is a steroid.

22. (Previously presented) The composition of Claim 21 wherein the steroid is cortisol, corticosterone, hydrocortisone, hydrocortisol, cortisone, prednisone, prednisolone, dexamethasone, beclomethasone, betamethasone, mometasone, mometasone furoate, budesonide, triamcinolone acetonide, or fluticasone.

23. (Previously presented) The composition of Claim 7 wherein the anti-inflammatory drug is a nonsteroidal, anti-inflammatory drug.

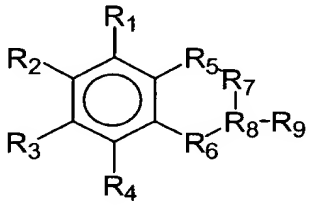
24. (Previously presented) The composition of Claim 23 wherein the nonsteroidal, anti-inflammatory drug is aspirin, acetominophen, ibuprofen, esculetin, phenidone, quercetin, ketoprofen, nordihydroguiaretic acid, sulindac, sulindac sulfone, sulindac sulfide, indomethacin, NS-398, cyclooxygenase-1 inhibitors, methylheptyl imidazole, furegrelate sodium, SKF525AHCL, thromboxane inhibitors, toradol, ecasa, salsalate, diflunisal, mefenamic acid, naproxen, naproxen sodium, flotafenine, meclofenamate, phenylbutazone, oxyphenbutazone, diclofenac, etodolac, fenoprofen, flufenamic acid, flurbiprofen, pirprofen, tolmetin, apazone, fenbufen, nabumetone, oxaprozin, piroxicam, salicylate, or tenoxicam.

25. (Previously presented) The composition of Claim 23 wherein the nonsteroidal, anti-inflammatory drug is indomethacin or sulindac.

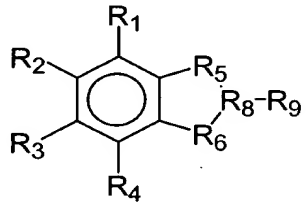
26. (Currently amended) A method for inhibiting angiogenesis in a human or animal comprising administering to a human or animal in need of such inhibition a composition comprising an angiogenesis inhibiting compound and an anti-inflammatory compound, wherein the angiogenesis inhibiting compound is:

(1) a compound the formula

A)

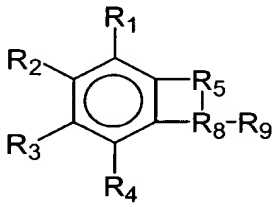


B)



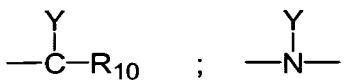
or

C)



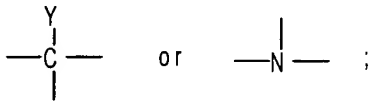
wherein

R_1-R_4 are each independently -H; -OH; =O; straight or branched chain alkane, alkene, or alkyne; cyclic alkane, alkene, or alkynes; a combinations of cyclic and acyclic alkanes, alkenes, or alkynes; alcohol, aldehyde, ketone, carboxylic acid, ester, or ether moieties in combination with acyclic, cyclic, or combination acyclic/cyclic moieties; aza; amino, $-XO_n$ or $-O-XO_n$, where X=N and n=2, X=S and n=2 or 3, or X=P and n=1-3; or halogen; R_5-R_7 are each independently



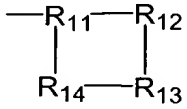
or -O-, where Y is absent and R_{10} is =O or Y and R_{10} are each independently -H; -OH; =O;

straight or branched chain alkane, alkene, or alkyne; cyclic alkane, alkene, or alkynes; a combination of cyclic and acyclic alkanes, alkenes, or alkynes; alcohol, aldehyde, ketone, carboxylic acid, ester, or ether moieties in combination with acyclic, cyclic, or combination acyclic/cyclic moieties; aza; amino, $-XO_n$ or $-O-XO_n$, where X=N and n=2, X=S and n=2 or 3, or X=P and n=1-3; or halogen; where R_8 is independently:

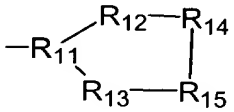


and R_9 is

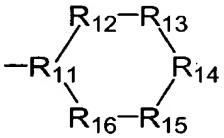
D)



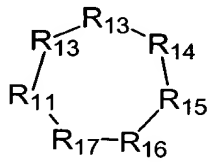
E)



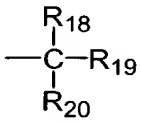
F)



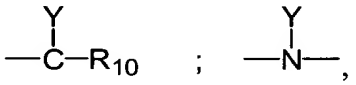
G)



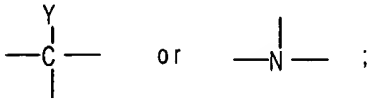
or H)



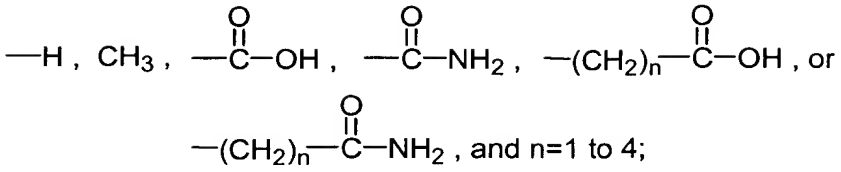
wherein each of R_{12}-R_{17} is independently



wherein R_{11} is

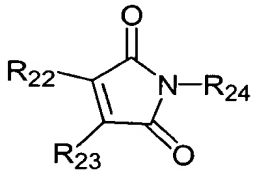


and wherein R_{18}, R_{19} and R_{20} are each independently



with the proviso that the angiogenesis inhibiting compound is not thalidomide;

(2) a compound of the formula

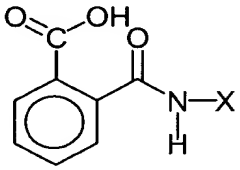


where R_{22} and R_{23} are each independently H, F, Cl, Br, I, CH_3, or $-CH_2-CH_3$;

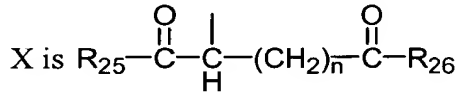
and R_{24} is H, CH_3, or $-CH_2-CH_3$;

or

(3) a compound of the formula



where X is R_6 as defined in (1) above, or

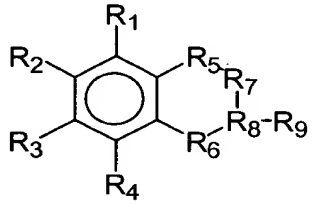


and R_{25} and R_{26} are independently -OH, -H, or $-NH_2$, and n = 1 through 4.

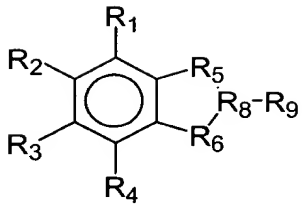
27. (Currently amended) A method for treating an angiogenesis dependent disease in a human or animal having such a disease comprising administering to the human or animal in need of such treatment a composition comprising an angiogenesis inhibiting compound and an anti-inflammatory compound, wherein the angiogenesis inhibiting compound is:

(1) a compound of the formula

A)

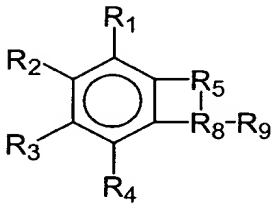


B)



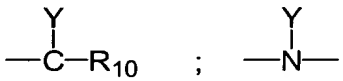
or

C)

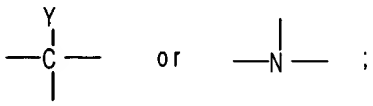


wherein

R_1-R_4 are each independently -H; -OH; =O; straight or branched chain alkane, alkene, or alkyne; cyclic alkane, alkene, or alkynes; a combinations of cyclic and acyclic alkanes, alkenes, or alkynes; alcohol, aldehyde, ketone, carboxylic acid, ester, or ether moieties in combination with acyclic, cyclic, or combination acyclic/cyclic moieties; aza; amino, $-XO_n$ or $-O-XO_n$, where X=N and n=2, X=S and n=2 or 3, or X=P and n=1-3; or halogen; R_5-R_7 are each independently

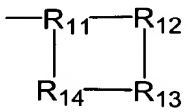


or -O-, where Y is absent and R_{10} is =O or Y and R_{10} are each independently -H; -OH; =O; straight or branched chain alkane, alkene, or alkyne; cyclic alkane, alkene, or alkynes; a combination of cyclic and acyclic alkanes, alkenes, or alkynes; alcohol, aldehyde, ketone, carboxylic acid, ester, or ether moieties in combination with acyclic, cyclic, or combination acyclic/cyclic moieties; aza; amino, $-XO_n$ or $-O-XO_n$, where X=N and n=2, X=S and n=2 or 3, or X=P and n=1-3; or halogen; where R_8 is independently:

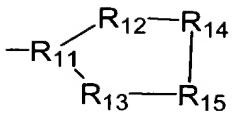


and R_9 is

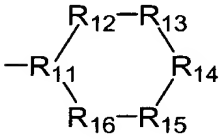
D)



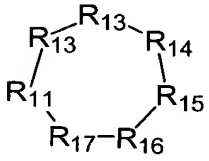
E)



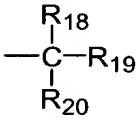
F)



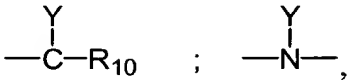
G)



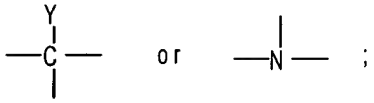
or H)



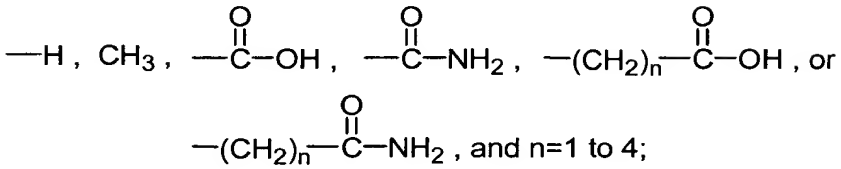
wherein each of R_{12}-R_{17} is independently



wherein R_{11} is

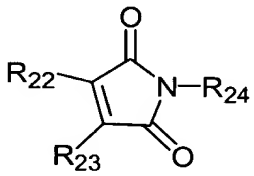


and wherein R_{18}, R_{19} and R_{20} are each independently



with the proviso that the angiogenesis inhibiting compound is not thalidomide;

(2) a compound of the formula

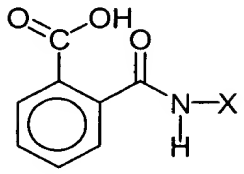


where R_{22} and R_{23} are each independently H, F, Cl, Br, I, CH_3, or $-CH_2-CH_3$;

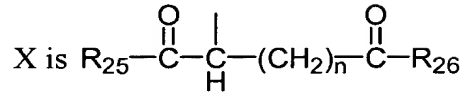
and R_{24} is H, CH_3, or $-CH_2-CH_3$;

or

(3) a compound of the formula



where X is R_6 as defined in (1) above, or



and R_{25} and R_{26} are independently -OH, -H, or $-NH_2$, and n = 1 through 4.

28. (Previously presented) The method of Claim 27 wherein the angiogenesis dependent disease is macular degeneration, diabetic retinopathy, neovascular glaucoma, retrolental fibroplasias, proliferative vitreoretinopathy, solid tumors, blood-borne tumors, leukemia, hemangioma, psoriasis, Kaposi's sarcoma, Chron's disease, ulcerative colitis, cancer, retinopathy of prematurity, corneal graft rejection, epidemic keratoconjunctivitis, Vitamin A deficiency, contact lens overwear, atopic keratitis, superior limbic keratitis, pterygium keratitis sicca, sjogren's syndrome, acne rosacea, phylectenulosis, syphilis, *Mycobacteria* infections, lipid degeneration, chemical burns, bacterial ulcers, fungal ulcers, *Herpes simplex* infections, *Herpes zoster* infections, Mooren's ulcer, Terrien's marginal degeneration, marginal keratolysis, trauma, rheumatoid arthritis systemic lupus, polyarteritis, Wegener's sarcoidosis, scleritis, Stevens-Johnson disease, radial keratotomy, corneal graft rejection, sickle cell anemia, pseudoxanthoma elasticum, pemphigoid, Paget's disease, vein occlusion, artery occlusion, carotid obstructive disease, chronic uveitis, chronic vitritis, Lyme's disease, systemic lupus erythematosis, Eales' disease, Behcet's disease, presumed ocular histoplasmosis, Best's disease, myopia, optic pits, Stargardt's disease, pars planitis, chronic retinal detachment, hyperviscosity syndromes, toxoplasmosis, post-laser complications, or rubeosis.

Claims 29-31 (Canceled).

32. (Previously presented) The method of Claim 26 wherein the composition is suitable for oral, rectal, ophthalmic, nasal, topical, vaginal, or parenteral administration.

33. (Currently amended) The method of Claim ~~22~~ 27 wherein the composition is suitable for oral, rectal, ophthalmic, nasal, topical, vaginal, or parenteral administration.

34. (Previously presented) The method of Claim 26 wherein the angiogenesis inhibiting compound is in a dosage of between about 0.1 mg/kg/day to about 300 mg/kg/day.

35. (Previously presented) The method of Claim 34 wherein the dosage of the angiogenesis inhibiting compound is between about 0.5 mg/kg/day to about 50 mg/kg/day.

36. (Previously presented) The method of Claim 35 wherein the dosage of the angiogenesis inhibiting compound is between about 1 mg/kg/day to about 10 mg/kg/day.

37. (Previously presented) The method of Claim 27 wherein the dosage of the angiogenesis inhibiting compound is between about 0.1 mg/kg/day to about 300 mg/kg/day.

38. (Previously presented) The method of Claim 37 wherein the dosage of the angiogenesis inhibiting compound is between about 0.5 mg/kg/day to about 50 mg/kg/day.

39. (Previously presented) The method of Claim 38 wherein the dosage of the angiogenesis inhibiting compound is between about 1 mg/kg/day to about 10 mg/kg/day.

40-42. (Canceled)

43. (Currently amended) ~~An~~ The ~~angiogenesis inhibitory~~ composition of Claim 11 wherein the anti-inflammatory drug is a nonsteroidal, anti-inflammatory drug.

44. (Previously presented) The composition of Claim 43 wherein the nonsteroidal, anti-inflammatory drug is aspirin, acetominophen, ibuprofen, esculetin, phenidone, quercetin, ketoprofen, nordihydroguiaretic acid, sulindac, sulindac sulfone,

sulindac sulfide, indomethacin, NS-398, cyclooxygenase-1 inhibitors, methylheptyl imidazole, furegrelate sodium, SKF525AHCL, thromboxane inhibitors, toradol, ecasa, salsalate, diflunisal, mefenamic acid, naproxen, naproxen sodium, flocafenine, meclofenamate, phenylbutazone, oxyphenbutazone, diclofenac, etodolac, fenoprofen, flufenamic acid, flurbiprofen, pirprofen, tolmetin, apazone, fenbufen, nabumetone, oxaprozin, piroxicam, salicylate, or tenoxicam.

45-46. (Canceled).

47. (Previously presented) The method of Claim 26, wherein the antiinflammatory compound is a nonsteroidal, antiinflammatory drug.

48. (Previously presented) The method of Claim 27, wherein the antiinflammatory compound is a nonsteroidal, antiinflammatory drug.

49. (New) The method of Claim 47, wherein the nonsteroidal, anti-inflammatory drug is aspirin, acetominophen, ibuprofen, esculetin, phenidone, quercetin, ketoprofen, nordihydroguiaretic acid, sulindac, sulindac sulfone, sulindac sulfide, indomethacin, NS-398, cyclooxygenase-1 inhibitors, methylheptyl imidazole, furegrelate sodium, SKF525AHCL, thromboxane inhibitors, toradol, ecasa, salsalate, diflunisal, mefenamic acid, naproxen, naproxen sodium, flotafenine, meclofenamate, phenylbutazone, oxyphenbutazone, diclofenac, etodolac, fenoprofen, flufenamic acid, flurbiprofen, pirprofen, tolmetin, apazone, fenbufen, nabumetone, oxaprozin, piroxicam, salicylate, or tenoxicam.

50. (New) The method of Claim 48, wherein the nonsteroidal, anti-inflammatory drug is aspirin, acetominophen, ibuprofen, esculetin, phenidone, quercetin, ketoprofen, nordihydroguiaretic acid, sulindac, sulindac sulfone, sulindac sulfide, indomethacin, NS-398, cyclooxygenase-1 inhibitors, methylheptyl imidazole, furegrelate sodium, SKF525AHCL, thromboxane inhibitors, toradol, ecasa, salsalate, diflunisal, mefenamic acid, naproxen, naproxen sodium, flotafenine, meclofenamate, phenylbutazone, oxyphenbutazone, diclofenac, etodolac, fenoprofen, flufenamic acid, flurbiprofen, pirprofen, tolmetin, apazone, fenbufen, nabumetone, oxaprozin, piroxicam, salicylate, or tenoxicam.

51. (New) The method of Claim 26 wherein the antiinflammatory drug is a steroid.

52. (New) The method of Claim 51 wherein the steroid is cortisol, corticosterone, hydrocortisone, hydrocortisol, cortisone, prednisone, prednisolone, dexamethasone, beclomethasone, betamethasone, mometasone, mometasone furoate, budesonide, triamcinolone acetonide, or fluticasone.

53. (New) The method of Claim 27 wherein antiinflammatory drug is a steroid.

54. (New) The method of Claim 53 wherein the steroid is cortisol, corticosterone, hydrocortisone, hydrocortisol, cortisone, prednisone, prednisolone, dexamethasone, beclomethasone, betamethasone, mometasone, mometasone furoate, budesonide, triamcinolone acetonide, or fluticasone.